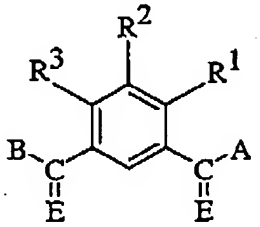
AMENDMENT TO THE CLAIMS

The following listing of claim(s) will replace all prior versions, and listings, of claim(s) in the application.

Listing of claim(s):

Claim 1 (amended). A method for inhibiting matrix metalloproteinase enzymes in a mammal comprising administering to the mammal an MMP inhibiting amount of a compound of Formula I



I

wherein:

R^1, R^2, and R^3 independently are hydrogen, halo, hydroxy, C_1-C_6 alkyl, C_1-C_6 alkoxy, C_2-C_6 alkenyl, C_2-C_6 alkynyl, NO_2, NR^4R^5, CN, or CF_3;

E is independently O or S;

A and B independently are OR^4 or NR^4R^5;

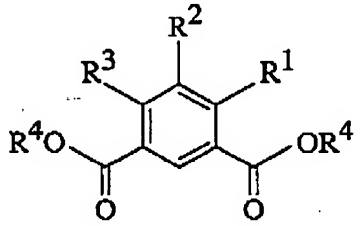
each R^4 and R^5 independently are ~~H, C_1-C_6 alkyl, C_2-C_6 alkenyl, C_2-C_6 alkynyl, $(CH_2)_n$ aryl, $(CH_2)_n$ cycloalkyl,~~ $(CH_2)_n$ heterocyclyl, $(CH_2)_n$ heteroaryl, or R^4 and R^5 when taken together with the nitrogen to which they are attached complete a 3- to 8-membered ring, optionally containing a heteroatom selected from O, S, or NH, and optionally substituted or unsubstituted;

n is an integer from 0 to 6;

or a pharmaceutically acceptable salt thereof;

wherein the compound isophthalic acid bis-(1,3-benzodioxol-5-ylmethyl) ester is excluded.

Claim 2 (amended). A method for inhibiting matrix metalloproteinase enzymes in a mammal comprising administering to the mammal an MMP inhibiting amount of a compound of Formula II



II

wherein:

R^1, R^2, and R^3 independently are hydrogen, halo, hydroxy, C_1-C_6 alkyl, C_1-C_6 alkoxy, C_2-C_6 alkenyl, C_2-C_6 alkynyl, NO_2, NR^4R^5, CN, or CF_3; and

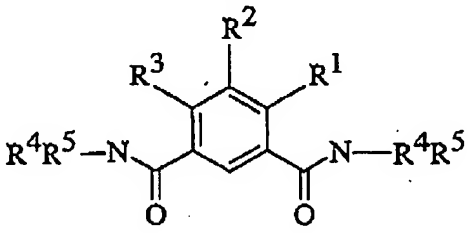
R^4 and R^5 is independently ~~H, C_1-C_6 alkyl, C_2-C_6 alkenyl, C_2-C_6 alkynyl, $(CH_2)_n$ aryl, $(CH_2)_n$ cycloalkyl, or~~ $(CH_2)_n$ heterocyclyl, $(CH_2)_n$ heteroaryl, or R^4 and R^5 when taken together with the nitrogen to which they are attached complete a 3- to 8-membered ring, optionally containing a heteroatom selected from O, S, or NH, and optionally substituted or unsubstituted;

n is an integer from 0 to 6;

or a pharmaceutically acceptable salt thereof;

wherein the compound isophthalic acid bis-(1,3-benzodioxol-5-ylmethyl) ester is excluded.

Claim 3 (amended). A method for inhibiting matrix metalloproteinase enzymes in a mammal comprising administering to the mammal an MMP inhibiting amount of a compound of Formula III



III

wherein:

R^1, R^2, and R^3 independently are hydrogen, halo, hydroxy, C_1-C_6 alkyl, C_1-C_6 alkoxy, C_2-C_6 alkenyl, C_2-C_6 alkynyl, NO_2, NR^4R^5, CN, or CF_3;

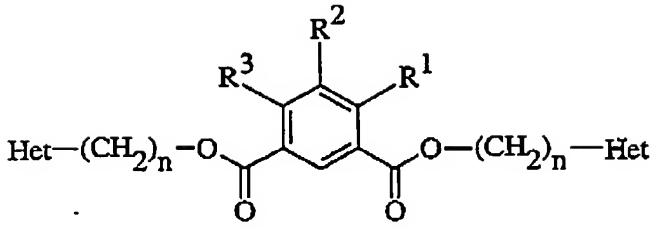
R^4 and R^5 independently are ~~H, C_1-C_6 alkyl, C_2-C_6 alkenyl, C_2-C_6 alkynyl, $(CH_2)_n$ aryl, $(CH_2)_n$ cycloalkyl,~~ $(CH_2)_n$ heterocyclyl, $(CH_2)_n$ heteroaryl, or R^4 and R^5 when taken together with the nitrogen to which they are attached complete a 3- to 8-membered ring, optionally containing a heteroatom selected from O, S, or NH, and optionally substituted or unsubstituted;

n is an integer from 0 to 6;

or a pharmaceutically acceptable salt thereof.

Claim 4 (canceled).

Claim 5 (amended). A method for inhibiting matrix metalloproteinase enzymes in a mammal comprising administering to the mammal an MMP inhibiting amount of a compound of Formula V



V

wherein:

R^1, R^2, and R^3 independently are hydrogen, halo, hydroxy, C_1-C_6 alkyl, C_1-C_6 alkoxy, C_2-C_6 alkenyl, C_2-C_6 alkynyl, NO_2, NR^4R^5, CN, or CF_3, and Het is an unsubstituted or substituted heteroaryl group;

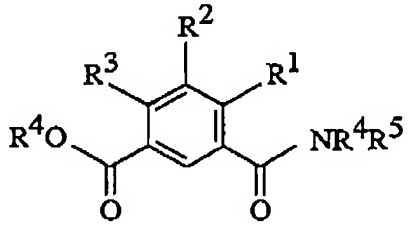
R^4 and R^5 independently are ~~H, C_1-C_6 alkyl, C_2-C_6 alkenyl, C_2-C_6 alkynyl, $(CH_2)_n$ aryl, $(CH_2)_n$ cycloalkyl,~~ $(CH_2)_n$ heterocyclyl, $(CH_2)_n$ heteroaryl, or R^4 and R^5 when taken together with the nitrogen to which they are attached complete a 3- to 8-membered ring, optionally containing a heteroatom selected from O, S, or NH, and optionally substituted or unsubstituted;

n is an integer from 0 to 6;

or a pharmaceutically acceptable salt thereof;

wherein the compound isophthalic acid bis-(1,3-benzodioxol-5-ylmethyl) ester is excluded.

Claim 6 (amended). A method for inhibiting matrix metalloproteinase enzymes in a mammal comprising administering to the mammal an MMP inhibiting amount of a compound of Formula VI



VI

or a pharmaceutically acceptable salt thereof,

wherein:

R^1, R^2, and R^3 independently are hydrogen, halo, hydroxy, C_1-C_6 alkyl, C_1-C_6 alkoxy, C_2-C_6 alkenyl, C_2-C_6 alkynyl, NO_2, NR^4R^5, CN, or CF_3;

R^4 and R^5 independently are ~~H, C_1-C_6 alkyl, C_2-C_6 alkenyl, C_2-C_6 alkynyl, $(CH_2)_n$ aryl, $(CH_2)_n$ cycloalkyl,~~ $\underline{(CH_2)_n}$ heterocyclyl, $(CH_2)_n$ heteroaryl, or R^4 and R^5 when taken together with the nitrogen to which they are attached complete a 3- to 8-membered ring, optionally containing a heteroatom selected from O, S, or NH, and optionally substituted or unsubstituted; and

n is an integer from 0 to 6.

Claim 7 (amended). A compound selected from:

~~4-Methoxy-N,N'-bis-(4-methoxybenzyl)-isophthalamide;~~

Isophthalic acid di-(2,1,3-benzothiadiazol-5-yl) methyl ester;

~~4-Methoxy-isophthalic-acid-dibenzyl-ester;~~

4-Methoxy-isophthalic acid dipyridin-4-ylmethyl ester;

~~Isophthalic-acid-bis-(4-fluoro-benzyl)-ester;~~

~~Isophthalic-acid-bis-(3-fluoro-benzyl)-ester;~~

~~Isophthalic-acid-bis-(4-methoxy-benzyl)-ester;~~

~~Isophthalic-acid-bis-(3-methoxy-benzyl)-ester;~~

~~Isophthalic-acid-bis-(1,3-benzodioxol-5-ylmethyl)-ester;~~

~~N,N'-Bis-(3-fluoro-benzyl)-isophthalamide;~~

~~4-Acetyl-isophthalic-acid-dibenzyl-ester;~~

~~4-Methoxycarbonylmethoxy-isophthalic-acid-dibenzyl-ester;~~

N,N'-Bis-1,3-benzodioxol-5-ylmethyl-4-methoxy-isophthalamide;

~~N-1,3-Benzodioxol-5-ylmethyl-4-methoxy-N'-(4-methoxy-benzyl)-isophthalamide;~~

~~4-Methoxy-N,N'-bis-(4-methoxy-benzyl)-isophthalamide;~~

~~N-1,3-Benzodioxol-5-ylmethyl N′-(4-chloro-benzyl)-4-methoxy-isophthalamide;~~

~~N-Benzyl-4-methoxy-N′-(4-methoxy-benzyl)-isophthalamide;~~

~~N′-Benzyl-4-methoxy-N-(4-methoxy-benzyl)-isophthalamide;~~

~~4-Methoxy-N-(4-methoxy-benzyl)-N′-pyridin-4-ylmethyl-isophthalamide;~~

~~N′-1,3-Benzodioxol-5-ylmethyl-4-methoxy-N-(2-phenoxy-ethyl)-isophthalamide;~~

~~N-1,3-Benzodioxol-5-ylmethyl-4-methoxy-N′-(2-phenoxy-ethyl)-isophthalamide;~~

N-1,3-Benzodioxol-5-ylmethyl-N′-furan-2-ylmethyl-isophthalamide;

~~N′-1,3-Benzodioxol-5-ylmethyl-N-(2-ethoxy-ethyl)-4-methoxy-isophthalamide;~~

~~N,N′-Bis-(3-hydroxymethyl-phenyl)-isophthalamide;~~

~~N-Benzyl-4-methoxy-N′-(2-phenoxy-ethyl)-isophthalamide;~~

~~4-Methoxy-N,N′-bis-(4-methyl-benzyl)-isophthalamide;~~

~~4-Methoxy-N,N′-bis-(3-methoxy-benzyl)-isophthalamide;~~

~~N-1,3-Benzodioxol-5-ylmethyl-4-methoxy-N′-(4-methoxy-benzyl)-isophthalamide;~~

~~N-1,3-Benzodioxol-5-ylmethyl-isophthalamic-acid,~~ ~~(4-carboxyphenyl)methyl-ester;~~

~~4-{[3-(3-Methoxy-benzylcarbamoyl)-benzoylamino]-methyl}-benzoic acid;~~

4-Methoxy-isophthalic acid di-2,1,3-benzothiadiazol-5-ylmethyl ester;

~~4-{[3-(3-Methoxy-benzylcarbamoyl)-benzoylamino]-methyl}-benzoic acid-methyl-ester;~~

~~N-(3-Methoxy-benzyl)-N′-(4-nitro-benzyl)-isophthalamide;~~

~~N-(3,4-Dichloro-benzyl)-N′-pyridin-4-ylmethyl-isophthalamide;~~

N1,N3-Bis-1,3-benzodioxol-5-ylmethyl-4-ethoxy-isophthalamide;

~~N-(4-Chloro-benzyl)-N′-(3-methoxy-benzyl)-isophthalamide;~~

~~N-(3,4-Dichloro-benzyl)-N'-(3-methoxy-benzyl)-isophthalamide;~~

~~N-(4-Methoxy-benzyl)-N'-(3-methoxy-benzyl)-isophthalamide;~~

~~N,N'-Bis-(4-fluoro-3-methoxy-benzyl)-isophthalamide;~~

~~4-Ethoxy-N1,N3-bis-(3-methoxy-benzyl)-isophthalamide;~~

N1,N3-Bis-1,3-benzodioxol-5-ylmethyl-4-ethoxy-isophthalamide;

~~N-(3-Methoxy-benzyl)-N'-pyridin-3-ylmethyl-isophthalamide;~~

~~N-(3-Methoxy-benzyl)-N'-pyridin-4-ylmethyl-isophthalamide;~~

N1-1,3-Benzodioxol-5-ylmethyl-N3-pyridin-3-ylmethyl-isophthalamide;

~~N-(3-Methoxy-benzyl)-N'-(3-trifluoromethoxy-benzyl)-isophthalamide;~~

N1,N3-Bis-1,3-benzodioxol-5-ylmethyl-4-isopropoxy-isophthalamide;

~~4-Isopropoxy-N1,N3-bis-(3-methoxy-benzyl)-isophthalamide;~~

~~N1-Benzyl-4-methoxy-N3-(4-methoxy-benzyl)-isophthalamide;~~

~~N1-1,3-Benzodioxol-5-ylmethyl-4-methoxy-N3-(4-methoxy-benzyl)-isophthalamide;~~

~~N1-1,3-Benzodioxol-5-ylmethyl-4-methoxy-N3-(2-phenoxy-ethyl)-isophthalamide;~~

~~N1-Benzyl-4-methoxy-N3-(2-phenoxy-ethyl)-isophthalamide;~~

~~N1-1,3-Benzodioxol-5-ylmethyl-N3-(4-chloro-benzyl)-4-methoxy-isophthalamide;~~

~~N3-1,3-Benzodioxol-5-ylmethyl-4-methoxy-N1-(4-methoxy-benzyl)-isophthalamide;~~

~~N3-Benzyl-4-methoxy-N1-(4-methoxy-benzyl)-isophthalamide;~~

~~N3-1,3-Benzodioxol-5-ylmethyl-4-methoxy-N1-(2-phenoxy-ethyl)-isophthalamide;~~

~~N3-1,3-Benzodioxol-5-ylmethyl-N1-(2-ethoxy-ethyl)-4-methoxy-isophthalamide;~~

~~4-Methoxy-N1-(4-methoxy-benzyl)-N3-pyridin-4-ylmethyl-isophthalamide;~~

4-Amino-N1,N3-bis-1,3-benzodioxol-5-ylmethyl-isophthalamide;

4-Acetylamino-N1,N3-bis-1,3-benzodioxol-5-ylmethyl-isophthalamide;

~~N (3-Methoxy-benzyl) N′-pyridin-3-ylmethyl-isophthalamide;~~

~~N (3-Methoxy-benzyl) N′-pyridin-4-ylmethyl-isophthalamide;~~

N1-1,3-Benzodioxol-5-ylmethyl-N3-pyridin-3-ylmethyl-isophthalamide;

~~N (4-Chloro-benzyl) N′-(3-methoxy-benzyl) isophthalamide;~~

~~N (3,4-Dichloro-benzyl) N′-(3-methoxy-benzyl) isophthalamide;~~

~~N (4-Methoxy-benzyl) N′-(3-methoxy-benzyl) isophthalamide;~~

~~N (3-Methoxy-benzyl) N′-(4-methyl-benzyl) isophthalamide;~~

~~N,N′-Bis (4-fluoro-3-methoxy-benzyl) isophthalamide;~~

~~({3-[(1,3-Benzodioxol-5-ylmethyl) carbamoyl] benzoyl}-benzyl-amino)-acetic acid;~~

~~N-Benzo[1,3]dioxol-5-ylmethyl-isophthalamic(4-hydroxymethyl-benzoic acid) ester;~~

~~N (3,4-Dichloro-benzyl) N′-pyridin-4-ylmethyl-isophthalamide;~~

~~N (3-Methoxy-benzyl) N′-(4-nitro-benzyl) isophthalamide;~~

~~4-{[3-(3-Methoxy-benzylcarbamoyl) benzoylamino]-methyl}-benzoic acid methyl ester;~~

~~N-3-methoxybenzyl-isophthalamic(4-hydroxymethyl-benzoic acid) ester;~~

~~4-{[3-(3-Methoxy-benzylcarbamoyl) benzoylamino]-methyl}-benzoic acid;~~

~~N (3-Amino-benzyl) N′-(3-methoxy-benzyl) isophthalamide;~~

~~N (3-Methoxy-benzyl) N′-(3-nitro-benzyl) isophthalamide;~~

~~4-Ethoxy-N′1,N″3-bis (3-methoxy-benzyl) isophthalamide;~~

N1,N3-Bis-1,3-benzodioxol-5-ylmethyl-4-ethoxy-isophthalamide;

N1,N3-Bis-1,3-benzodioxol-5-ylmethyl-4-propoxy-isophthalamide;

N1,N3-Bis-1,3-benzodioxol-5-ylmethyl-4-isopropoxy-isophthalamide;

N1,N3-Bis-2,1,3-benzothiadiazol-5-ylmethyl-4-methoxy-isophthalamide; and

4-Methoxy-isophthalic acid di-2,1,3-benzothiadiazol-5-ylmethyl ester.

Claim 8 (original). A pharmaceutical composition, comprising a compound of Claim 1, or a pharmaceutically acceptable salt thereof, admixed with a pharmaceutically acceptable carrier, diluent, or excipient.

Claim 9 (original). A pharmaceutical composition for inhibiting MMP-13 in a mammal, comprising an MMP-13 inhibiting amount of a compound of Claim 1, or a pharmaceutically acceptable salt thereof, admixed with a pharmaceutically acceptable carrier, diluent, or excipient.

Claim 10 (amended). A method for inhibiting MMP-13 in an animal, comprising administering to the animal an MMP-13 inhibiting amount of a compound of ~~Formula I~~ Claim 1, or a pharmaceutically acceptable salt thereof.

Claims 11 and 12 (canceled).

Claim 13 (amended).. A method for treating breast carcinoma, comprising administering to a patient suffering from such a disease an anticancer effective amount of a compound of ~~Formula I~~ Claim 1, or a pharmaceutically acceptable salt thereof.

Claim 14 (amended). A method for treating a rheumatoid arthritis, comprising administering to a patient suffering from such a disease an effective amount of a compound of ~~Formula I~~ Claim 1, or a pharmaceutically acceptable salt thereof.

Claim 15 (amended). A method for treating a osteoarthritis, comprising administering to a patient suffering from such a disease an effective amount of a compound of ~~Formula I~~ Claim 1, or a pharmaceutically acceptable salt thereof.

Claim 16 (amended). A method for treating a heart failure, comprising administering to a patient suffering from such a disease an effective amount of a compound of ~~Formula I~~ Claim 1, or a pharmaceutically acceptable salt thereof.

Claim 17 (amended). A method for treating a inflammation, comprising administering to a patient suffering from such a disease an effective amount of a compound of ~~Formula I~~ Claim 1, or a pharmaceutically acceptable salt thereof.